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NEWS 6 MAY 19 Derwent World Patents Index to be reloaded and enhanced  
NEWS 7 MAY 30 IPC 8 Rolled-up Core codes added to CA/CAPLUS and  
USPATFULL/USPAT2  
NEWS 8 MAY 30 The F-Term thesaurus is now available in CA/CAPLUS  
NEWS 9 JUN 02 The first reclassification of IPC codes now complete in  
INPADOC  
NEWS 10 JUN 26 TULSA/TULSA2 reloaded and enhanced with new search and  
and display fields  
NEWS 11 JUN 28 Price changes in full-text patent databases EPFULL and PCTFULL  
NEWS 12 JUL 11 CHEMSAFE reloaded and enhanced  
NEWS 13 JUL 14 FSTA enhanced with Japanese patents  
NEWS 14 JUL 19 Coverage of Research Disclosure reinstated in DWPI  
NEWS 15 AUG 09 INSPEC enhanced with 1898-1968 archive  
NEWS 16 AUG 28 ADISCTI Reloaded and Enhanced  
NEWS 17 AUG 30 CA(SM)/CAPLUS(SM) Austrian patent law changes  
NEWS 18 SEP 11 CA/CAPLUS enhanced with more pre-1907 records  
  
NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT  
MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.  
  
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FILE 'HOME' ENTERED AT 15:06:58 ON 20 SEP 2006

=> file caplus

COST IN U.S. DOLLARS

FULL ESTIMATED COST

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ENTRY

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FILE 'CAPLUS' ENTERED AT 15:07:09 ON 20 SEP 2006  
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FILE LAST UPDATED: 19 Sep 2006 (20060919/ED)

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=> s HAUSP and MDM2  
51 HAUSP  
2970 MDM2  
L1 21 HAUSP AND MDM2

=> s l1 not py>2004  
2147267 PY>2004  
L2 6 L1 NOT PY>2004

=> d ibib 1-6

L2 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2005:63694 CAPLUS  
DOCUMENT NUMBER: 143:224203  
TITLE: Dynamics in the p53-Mdm2 ubiquitination pathway  
AUTHOR(S): Brooks, Christopher L.; Gu, Wei  
CORPORATE SOURCE: Institute for Cancer Genetics and Department of Pathology; College of Physicians and Surgeons, Columbia University, New York, NY, USA  
SOURCE: Cell Cycle (2004), 3(7), 895-899  
CODEN: CCEYAS; ISSN: 1538-4101  
PUBLISHER: Landes Bioscience  
DOCUMENT TYPE: Journal; General Review  
LANGUAGE: English  
REFERENCE COUNT: 54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2005:60430 CAPLUS  
DOCUMENT NUMBER: 142:215611  
TITLE: HAUSP is required for p53 destabilization  
AUTHOR(S): Cummins, Jordan M.; Vogelstein, Bert  
CORPORATE SOURCE: The Howard Hughes Medical Institute, The Sidney Kimmel Comprehensive Cancer Center, Program in Cellular and Molecular Medicine, The Johns Hopkins University Medical Institutions, Baltimore, MD, USA  
SOURCE: Cell Cycle (2004), 3(6), 689-692  
CODEN: CCEYAS; ISSN: 1538-4101

PUBLISHER: Landes Bioscience  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:900604 CAPLUS  
DOCUMENT NUMBER: 142:4278  
TITLE: HAUSP/USP7 as an Epstein-Barr virus target  
AUTHOR(S): Holowaty, M. N.; Frappier, L.  
CORPORATE SOURCE: Department of Medical Genetics and Microbiology,  
University of Toronto, Toronto, Can.  
SOURCE: Biochemical Society Transactions (2004), 32(5),  
731-732  
CODEN: BCSTB5; ISSN: 0300-5127  
PUBLISHER: Portland Press Ltd.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:398363 CAPLUS  
DOCUMENT NUMBER: 141:121361  
TITLE: P53 apoptotic pathway molecules are frequently and  
simultaneously altered in nonsmall cell lung carcinoma  
AUTHOR(S): Mori, Shoichi; Ito, Genshi; Usami, Noriyasu; Yoshioka,  
Hiromu; Ueda, Yuichi; Kodama, Yoshinori; Takahashi,  
Masahide; Fong, Kwun M.; Shimokata, Kaoru; Sekido,  
Yoshitaka  
CORPORATE SOURCE: Department of Clinical Preventive Medicine, Department  
of Thoracic Surgery, Nagoya University School of  
Medicine, Nagoya, Japan  
SOURCE: Cancer (New York, NY, United States) (2004), 100(8),  
1673-1682  
CODEN: CANCAR; ISSN: 0008-543X  
PUBLISHER: John Wiley & Sons, Inc.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
REFERENCE COUNT: 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:312009 CAPLUS  
DOCUMENT NUMBER: 140:300911  
TITLE: A dynamic role of HAUSP in the p53-  
Mdm2 pathway  
AUTHOR(S): Li, Muyang; Brooks, Christopher L.; Kon, Ning; Gu, Wei  
CORPORATE SOURCE: Institute for Cancer Genetics and Department of  
Pathology College of Physicians and Surgeons, Columbia  
University, New York, NY, 10032, USA  
SOURCE: Molecular Cell (2004), 13(6), 879-886  
CODEN: MOCEFL; ISSN: 1097-2765  
PUBLISHER: Cell Press  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
REFERENCE COUNT: 34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2002:312567 CAPLUS  
DOCUMENT NUMBER: 137:44608  
TITLE: Deubiquitination of p53 by HAUSP is an

important pathway for p53 stabilization

AUTHOR(S): Li, Muyang; Chen, Delin; Shiloh, Ariel; Luo, Jianyuan; Nikolaev, Anatoly Y.; Qin, Jun; Gu, Wei

CORPORATE SOURCE: Institute for Cancer Genetics, and Department of Pathology, College of Physicians b Surgeons, Columbia University, New York, NY, 10032, USA

SOURCE: Nature (London, United Kingdom) (2002), 416(6881), 648-652

CODEN: NATUAS; ISSN: 0028-0836

PUBLISHER: Nature Publishing Group

DOCUMENT TYPE: Journal

LANGUAGE: English

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> s usp7  
L3 40 USP7

=> s l3 and MDM2  
2970 MDM2  
L4 8 L3 AND MDM2

=> d ibib 1-8

L4 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:197641 CAPLUS

DOCUMENT NUMBER: 144:288171

TITLE: Molecular recognition of p53 and MDM2 by USP7/HAUSP

AUTHOR(S): Sheng, Yi; Saridakis, Vivian; Sarkari, Feroz; Duan, Shili; Wu, Tianne; Arrowsmith, Cheryl H.; Frappier, Lori

CORPORATE SOURCE: Department of Medical Biophysics, Ontario Cancer Institute, Toronto, ON, M5G 1L7, Can.

SOURCE: Nature Structural & Molecular Biology (2006), 13(3), 285-291

CODEN: NSMBCU; ISSN: 1545-9993

PUBLISHER: Nature Publishing Group

DOCUMENT TYPE: Journal

LANGUAGE: English

REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:156572 CAPLUS

DOCUMENT NUMBER: 145:119254

TITLE: Structural basis of competitive recognition of p53 and MDM2 by HAUSP/USP7: implications for the regulation of the p53-MDM2 pathway

AUTHOR(S): Hu, Min; Gu, Lichuan; Li, Muyang; Jeffrey, Philip D.; Gu, Wei; Shi, Yigong

CORPORATE SOURCE: Department of Molecular Biology, Lewis Thomas Laboratory, Princeton University, Princeton, NJ, USA

SOURCE: PLoS Biology (2006), 4(2), 228-239

CODEN: PBLIBG; ISSN: 1545-7885

URL: [http://biology.plosjournals.org/archive/1545-7885/4/2/pdf/10.1371\\_1545-7885\\_4\\_2\\_complete.pdf](http://biology.plosjournals.org/archive/1545-7885/4/2/pdf/10.1371_1545-7885_4_2_complete.pdf)

PUBLISHER: Public Library of Science

DOCUMENT TYPE: Journal; (online computer file)

LANGUAGE: English

REFERENCE COUNT: 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:1056192 CAPLUS  
 DOCUMENT NUMBER: 143:455700  
 TITLE: Reciprocal activities between herpes simplex virus type 1 regulatory protein ICP0, a ubiquitin E3 ligase, and ubiquitin-specific protease USP7  
 AUTHOR(S): Boutell, Chris; Canning, Mary; Orr, Anne; Everett, Roger D.  
 CORPORATE SOURCE: MRC Virology Unit, Institute of Virology, Glasgow, G11 5JR, UK  
 SOURCE: Journal of Virology (2005), 79(19), 12342-12354  
 CODEN: JOVIAM; ISSN: 0022-538X  
 PUBLISHER: American Society for Microbiology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 REFERENCE COUNT: 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:327153 CAPLUS  
 DOCUMENT NUMBER: 143:2872  
 TITLE: Structure of the p53 binding domain of HAUSP/USP7 bound to Epstein-Barr nuclear antigen 1: Implications for EBV-mediated immortalization  
 AUTHOR(S): Saridakis, Vivian; Sheng, Yi; Sarkari, Feroz; Holowaty, Melissa N.; Shire, Kathy; Nguyen, Tin; Zhang, Rongguang G.; Liao, Jack; Lee, Weontae; Edwards, Aled M.; Arrowsmith, Cheryl H.; Frappier, Lori  
 CORPORATE SOURCE: Department of Medical Genetics and Microbiology, University of Toronto, Toronto, ON, M5S 1A8, Can.  
 SOURCE: Molecular Cell (2005), 18(1), 25-36  
 CODEN: MOCEFL; ISSN: 1097-2765  
 PUBLISHER: Cell Press  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:60430 CAPLUS  
 DOCUMENT NUMBER: 142:215611  
 TITLE: HAUSP is required for p53 destabilization  
 AUTHOR(S): Cummins, Jordan M.; Vogelstein, Bert  
 CORPORATE SOURCE: The Howard Hughes Medical Institute, The Sidney Kimmel Comprehensive Cancer Center, Program in Cellular and Molecular Medicine, The Johns Hopkins University Medical Institutions, Baltimore, MD, USA  
 SOURCE: Cell Cycle (2004), 3(6), 689-692  
 CODEN: CCEYAS; ISSN: 1538-4101  
 PUBLISHER: Landes Bioscience  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:1997 CAPLUS  
 DOCUMENT NUMBER: 142:111841  
 TITLE: Gene expression profiles and biomarkers for the detection of depression-related and other disease-related gene transcripts in blood  
 INVENTOR(S): Liew, Choong-Chin  
 PATENT ASSIGNEE(S): Chondrogene Limited, Can.

SOURCE: U.S. Pat. Appl. Publ., 154 pp., Cont.-in-part of U.S.  
Ser. No. 802,875.  
CODEN: USXXCO

DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 31  
PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO. | DATE        |
|------------------------|------|----------|-----------------|-------------|
| US 2004265868          | A1   | 20041230 | US 2004-812702  | 20040330    |
| US 2004014059          | A1   | 20040122 | US 2002-268730  | 20021009    |
| US 2006134635          | A1   | 20060622 | US 2004-802875  | 20040312    |
| US 2005191637          | A1   | 20050901 | US 2004-803737  | 20040318    |
| US 2005196762          | A1   | 20050908 | US 2004-803759  | 20040318    |
| US 2005196763          | A1   | 20050908 | US 2004-803857  | 20040318    |
| US 2005196764          | A1   | 20050908 | US 2004-803858  | 20040318    |
| US 2005208505          | A1   | 20050922 | US 2004-803648  | 20040318    |
| PRIORITY APPLN. INFO.: |      |          | US 1999-115125P | P 19990106  |
|                        |      |          | US 2000-477148  | B1 20000104 |
|                        |      |          | US 2002-268730  | A2 20021009 |
|                        |      |          | US 2003-601518  | A2 20030620 |
|                        |      |          | US 2004-802875  | A2 20040312 |
|                        |      |          | US 2001-271955P | P 20010228  |
|                        |      |          | US 2001-275017P | P 20010312  |
|                        |      |          | US 2001-305340P | P 20010713  |
|                        |      |          | US 2002-85783   | A2 20020228 |

L4 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:900604 CAPLUS

DOCUMENT NUMBER: 142:4278

TITLE: HAUSP/USP7 as an Epstein-Barr virus target

AUTHOR(S): Holowaty, M. N.; Frappier, L.

CORPORATE SOURCE: Department of Medical Genetics and Microbiology,  
University of Toronto, Toronto, Can.

SOURCE: Biochemical Society Transactions (2004), 32(5),  
731-732  
CODEN: BCSTB5; ISSN: 0300-5127

PUBLISHER: Portland Press Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:114335 CAPLUS

DOCUMENT NUMBER: 132:332744

TITLE: A genome-wide survey of RAS transformation targets

AUTHOR(S): Zuber, Johannes; Tchernitsa, Oleg I.; Hinzmann, Bernd;  
Schmitz, Anne-Chantal; Grips, Martin; Hellriegel,  
Martin; Sers, Christine; Rosenthal, Andre; Schafer,  
Reinhold

CORPORATE SOURCE: Laboratory of Molecular Tumour Pathology, Institute of  
Pathology, Charite, Humboldt-University, Berlin,  
D-10117, Germany

SOURCE: Nature Genetics (2000), 24(2), 144-152  
CODEN: NGENEC; ISSN: 1061-4036

PUBLISHER: Nature America

DOCUMENT TYPE: Journal

LANGUAGE: English

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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MOST RECENT UPDATE WEEK: 200637 <200637/EW>  
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>>> IMAGES ARE AVAILABLE ONLINE AND FOR EMAIL-PRINTS <<<

>>> NEW IPC8 DATA AND FUNCTIONALITY NOW AVAILABLE IN THIS FILE.

SEE

<http://www.stn-international.de/stndatabases/details/ipc-reform.html> >>>

>>> FOR CHANGES IN PCTFULL PLEASE SEE HELP CHANGE  
(last updated April 10, 2006) <<<

>>> NEW PRICES IN PCTFULL AS OF 01 JULY 2006. FOR DETAILS,  
PLEASE SEE HELP COST <<<

=> s USP7

L5 37 USP7

=> s HAUSP

L6 34 HAUSP

=> s 16 or 15

L7 59 L6 OR L5

=> s MDM2 and 17

829 MDM2

L8 18 MDM2 AND L7

=> s screen? or ident?

194428 SCREEN?

478664 IDENT?

L9 532010 SCREEN? OR IDENT?

=> s 19 and 18

L10 18 L9 AND L8

=> s 110 not py>2002

444636 PY>2002

L11 5 L10 NOT PY>2002

=> d ibib 1-5

L11 ANSWER 1 OF 5

ACCESSION NUMBER:

TITLE (ENGLISH):

TITLE (FRENCH):

INVENTOR(S):

PATENT ASSIGNEE(S):

PCTFULL COPYRIGHT 2006 Univentio on STN

2002070742 PCTFULL ED 20020926 EW 200237

METHOD FOR THE DEVELOPMENT OF GENE PANELS FOR  
DIAGNOSTIC AND THERAPEUTIC PURPOSES BASED ON THE  
EXPRESSION AND METHYLATION STATUS OF THE GENES  
PROCEDE DE MISE AU POINT DE GROUPES D'ECHANTILLONS DE  
GENES A DES FINS DE DIAGNOSTIC ET DE THERAPIE QUI SONT  
BASES SUR L'EXPRESSION ET L'ETAT DE METHYLATION DES  
GENES

OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
DE;

BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahndorf, DE

EPIGENOMICS AG, Kastanienallee 24, 10435 Berlin, DE [DE,

AGENT: DE]  
 SCHOHE, Stefan\$, Boehmert & Boehmert, Pettenkoferstr.  
 20-22, 80336 Muenchen\$, DE  
 LANGUAGE OF FILING: English  
 LANGUAGE OF PUBL.: English  
 DOCUMENT TYPE: Patent  
 PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002070742 | A1   | 20020912 |

DESIGNATED STATES

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
 CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
 SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW  
 RW (ARIPO): GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW  
 RW (EAPO): AM AZ BY KG KZ MD RU TJ TM  
 RW (EPO): AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR  
 RW (OAPI): BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG  
 APPLICATION INFO.: WO 2002-EP2255 A 20020301  
 PRIORITY INFO.: US 2001-60/272,549 20010301

L11 ANSWER 2 OF 5  
 ACCESSION NUMBER:  
 TITLE (ENGLISH):

PCTFULL COPYRIGHT 2006 Univentio on STN  
 2002070741 PCTFULL ED 20020926 EW 200237  
 METHODS, SYSTEMS AND COMPUTER PROGRAM PRODUCTS FOR  
 DETERMINING THE BIOLOGICAL EFFECT AND/OR ACTIVITY OF  
 DRUGS, CHEMICAL SUBSTANCES AND/OR PHARMACEUTICAL  
 COMPOSITIONS BASED ON THEIR EFFECT ON THE METHYLATION  
 STATUS OF THE DNA

TITLE (FRENCH):

PROCEDES, SYSTEMES ET PRODUITS PROGRAMMES INFORMATIQUES  
 PERMETTANT DE DETERMINER L'EFFET BIOLOGIQUE ET/OU  
 L'ACTIVITE DE MEDICAMENTS, DE SUBSTANCES CHIMIQUES  
 ET/OU DE COMPOSITIONS PHARMACEUTIQUES, SUR LA BASE DE  
 LEUR EFFET SUR L'ETAT DE METHYLATION DE L'ADN

INVENTOR(S):

OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
 DE;

PATENT ASSIGNEE(S):

BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahnsdorf, DE  
 EPIGENOMICS AG, Kastanienallee 24, 10435 Berlin, DE  
 [DE, DE]

AGENT:

SCHOHE, Stefan\$, Boehmert & Boehmert,  
 Pettenkoferstrasse 20-22, 80336 Muenchen\$, DE

LANGUAGE OF FILING:

English

LANGUAGE OF PUBL.:

English

DOCUMENT TYPE:

Patent

PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002070741 | A2   | 20020912 |

DESIGNATED STATES

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 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
 SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW  
 RW (ARIPO): GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW  
 RW (EAPO): AM AZ BY KG KZ MD RU TJ TM  
 RW (EPO): AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR  
 RW (OAPI): BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG  
 APPLICATION INFO.: WO 2002-EP2254 A 20020301  
 PRIORITY INFO.: US 2001-60/272,484 20010301



L11 ANSWER 3 OF 5  
ACCESSION NUMBER:  
TITLE (ENGLISH):  
TITLE (FRENCH):  
INVENTOR(S):

PCTFULL COPYRIGHT 2006 Univentio on STN  
2002057414 PCTFULL ED 20020801 EW 200230  
LEUKOCYTE EXPRESSION PROFILING  
EVALUATION DU NIVEAU D'EXPRESSION LEUCOCYTAIRE  
WOHLGEMUTH, Jay, 664 Hamilton Avenue, Palo Alto, CA  
94301, US [US, US];  
FRY, Kirk, 2604 Ross Road, Palo Alto, CA 94303, US [US,  
US];  
MATCUK, George, 141C Escondido Village, Stanford, CA  
94305, US [US, US];  
ALTMAN, Peter, 717 Evelyn Avenue, Albany, CA 94706, US  
[US, US];  
PRENTICE, James, 120 Dolores Street, San Francisco, CA  
94103, US [US, US];  
PHILLIPS, Julie, 1090 Mirador Terrace, Pacifica, CA  
94044, US [US, US];  
LY, Ngoc, 2000 Crystal Springs Road 15-14, San Bruno,  
CA 94066, US [US, US];  
WOODWARD, Robert, 1828 Rheem Court, Pleasanton, CA  
94588, US [US, US];  
QUENTERMOUS, Thomas, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US];  
JOHNSON, Frances, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US]

PATENT ASSIGNEE(S):

BIOCARDIA, INC., 384 Oyster Point Boulevard, #4, South  
San Francisco, CA 94080, US [US, US], for all  
designates States except US;  
WOHLGEMUTH, Jay, 664 Hamilton Avenue, Palo Alto, CA  
94301, US [US, US], for US only;  
FRY, Kirk, 2604 Ross Road, Palo Alto, CA 94303, US [US,  
US], for US only;  
MATCUK, George, 141C Escondido Village, Stanford, CA  
94305, US [US, US], for US only;  
ALTMAN, Peter, 717 Evelyn Avenue, Albany, CA 94706, US  
[US, US], for US only;  
PRENTICE, James, 120 Dolores Street, San Francisco, CA  
94103, US [US, US], for US only;  
PHILLIPS, Julie, 1090 Mirador Terrace, Pacifica, CA  
94044, US [US, US], for US only;  
LY, Ngoc, 2000 Crystal Springs Road 15-14, San Bruno,  
CA 94066, US [US, US], for US only;  
WOODWARD, Robert, 1828 Rheem Court, Pleasanton, CA  
94588, US [US, US], for US only;  
QUENTERMOUS, Thomas, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US], for US only;  
JOHNSON, Frances, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US], for US only  
WARD, Michael, R.\$, Morrison & Foerster LLP, 425 Market  
Street, San Francisco, CA 94105-2482\$, US

AGENT:

LANGUAGE OF FILING:  
LANGUAGE OF PUBL.:  
DOCUMENT TYPE:  
PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002057414 | A2   | 20020725 |

DESIGNATED STATES

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CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
MG MK MN MW MX MZ NO NZ PH PL PT RO RU SD SE SG SI SK  
SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW  
GH GM KE LS MW MZ SD SL SZ TZ UG ZW  
AM AZ BY KG KZ MD RU TJ TM

RW (ARIPO):

RW (EAPO):

RW (EPO): AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
TR  
RW (OAPI): BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG  
APPLICATION INFO.: WO 2001-US47856 A 20011022  
PRIORITY INFO.: US 2000-60/241,994 20001020  
US 2001-60/296,764 20010608

L11 ANSWER 4 OF 5 PCTFULL COPYRIGHT 2006 Univentio on STN  
ACCESSION NUMBER: 2000079267 PCTFULL ED 20020515  
TITLE (ENGLISH): TREATMENT OF CANCER  
TITLE (FRENCH): TRAITEMENT ANTICANCEREUX  
INVENTOR(S): NIZETIC, Dean;  
GROET, JuergenRP : GILL JENNINGS & EVERY  
PATENT ASSIGNEE(S): SCHOOL OF PHARMACY, UNIVERSITY OF LONDON;  
NIZETIC, Dean;  
GROET, Juergen  
LANGUAGE OF PUBL.: English  
DOCUMENT TYPE: Patent  
PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2000079267 | A2   | 20001228 |

DESIGNATED STATES  
W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU  
CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN  
IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK  
MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM  
TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD  
SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY  
DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ CF CG  
CI CM GA GN GW ML MR NE SN TD TG  
WO 2000-GB2446 A 20000622  
GB 2000-0008161.2 20000403  
GB 1999-9914589.8 19990622

APPLICATION INFO.:  
PRIORITY INFO.:

L11 ANSWER 5 OF 5 PCTFULL COPYRIGHT 2006 Univentio on STN  
ACCESSION NUMBER: 2000073479 PCTFULL ED 20020515  
TITLE (ENGLISH): A COMBINED GROWTH FACTOR-DELETED AND THYMIDINE  
KINASE-DELETED VACCINIA VIRUS VECTOR  
TITLE (FRENCH): VECTEUR DU VIRUS DE LA VACCINE COMBINANT DELETION DU  
FACTEUR DE CROISSANCE ET DELETION DE THYMIDINE KINASE  
INVENTOR(S): MCCART, J., Andrea;  
BARTLETT, David, L.;  
MOSS, BernardRP : NATAUPSKY, Steven, J.  
PATENT ASSIGNEE(S): THE GOVERNMENT OF THE UNITED STATES OF AMERICA, as  
represented by THE SECRETARY, DEPARTMENT OF HEALTH AND  
HUMAN SERVICES;  
MCCART, J., Andrea;  
BARTLETT, David, L.;  
MOSS, Bernard  
LANGUAGE OF PUBL.: English  
DOCUMENT TYPE: Patent  
PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2000073479 | A1   | 20001207 |

DESIGNATED STATES  
W:

AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ  
DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS  
JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN  
MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT  
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TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY DE DK  
ES FI FR GB GR IE IT LU MC NL PT SE BF BJ CF CG CI CM

|                    |                            |            |
|--------------------|----------------------------|------------|
|                    | GA GN GW ML MR NE SN TD TG |            |
| APPLICATION INFO.: | WO 2000-US14679            | A 20000526 |
| PRIORITY INFO.:    | US 1999-60/137,126         | 19990528   |

=> d kwic 4

L11 ANSWER 4 OF 5 PCTFULL COPYRIGHT 2006 Univentio on STN

DETD . . . as p53, could perhaps explain the link to deletions of USPs in solid tumours. De-ubiquitination could play a major role in the Mdm2 mediated control of p53 levels and its activation mechanism, since the ubiquitin-mediated proteasome degradation of p53 is an important effector arm of. . .

In recent years a number of other protein modifying polypeptide tags have been identified. Many of these are related to ubiquitin and-have high levels of identity and similarity (determined using the BLAST algorithm, for instance) to ubiquitin itself. There is a recognised super family of such proteins which. . .

human SUMO-1 (PIC1 1 Sentrin, hSmt3C), SUMO-2 (hSmt3A) and SUMO-3 (hSMT3B) belong to the same family of UbL proteins with approximately 50% identity between themselves, and some 15-30% identity and 40-60% similarity in amino acid sequence to ubiquitin (Lapenta et al. 1997, Mannen et al. 1996, Kamitani et al. 1998, Saitoh. . .

Several UbL hydrolase enzymes have been identified which convert precursor UbL to active UbL. Some such enzymes interact with ubiquitin itself as well as with other UbL's. Proteases involved in cleavage of conjugates of UbL with target protein have been identified for instance SENP1 and SUSP-1, which were recently cloned (Kim et al. 2000, Gong et al. 2000a), and found to specifically cleave. . .

Valero, et al. (1 999) published after the first priority date of the present application, have, in parallel identified this gene and pointed out the gene product's sequence homologies to known USIP's in the conserved peptide domains previously identified e.g. by d'Andrea et al (1 998). They postulate a role in Alzheimer's disease. This protein has the HUGO approved name USP25.

fusion protein of the ubiquitin-like protein of interest and a detectable protein, and using the usual separation and immune based or autoradiographic identification techniques.

the specified domains, some level of sequence homology with sequence ID 1 , for instance at least

20%, preferably at least 50%, identity with that sequence, and a level of similarity of at least 50%, preferably at least 70% or more with that sequence (in. . .

the corresponding mouse product, described in Valero, et al 1999 may be used or sequences which have the above levels of identity and similarity with such a sequence.

outside the specified domains, some level of sequence homology with sequence ID 1 for instance at least 20%, preferably at least 50%, identity with that sequence, and a level of similarity of at least 50%, preferably at least 70% or more with that sequence (in. . . as the corresponding mouse product, described in Valero, et al 1999 may be used or sequences which have the above levels of identity and similarity with such a sequence.

#### Experimental

We identify a portion of human chromosome 21 homozygously deleted in non-small cell non carcinoma (NSCLC) for further study. The region contained the DNA. . . et al. We found a shared region of overlap (SRO) for the hemizygous loss in other NSCLC. The current work is to identify genes in the SRO which have a potential role in tumour suppression.

The exposure was for 14 hr to Molecular Dynamics (Sunnyvale, CA) Phosphorimager screens. The I.M.A.G.E. Consortium (Lennon et al., 1996) cDNA clone ID 82471 0 and the Unigene clone A0021343 have been used as labelled. . .

#### Identification and cloning of USP26

Twelve sequenced exon-trapped products, when analysed using BLAST-N against public sequence databases, revealed clusters of overlapping cDNA clones. Sequences. . .

with the binding of USP25 to its natural ubiquitinated substrates, since this residue is conserved between all UCH-s and USP-s so far identified.

of the sequences found to be interacting, from the GenBank database are given in the table, Table 1. Summary of frequency and identities of specific interacting proteins from human brain with USP25-C178A, detected using Yeast-Two-Hybrid technique  
Summary of Results by Number of specific Accession number decreasing. . .

#### SUMO-

1 (PIC1, Sentrin, hSmt3C), SUMO-2 (hSmt3A) and SUMO-3 (hSMT3B) belong to the same family of UbL proteins with approximately 50% identity between themselves, and some 15-30% identity and 40-60% similarity in

amino acid sequence to ubiquitin (Lapenta et al. 1997, Mannen et al. 1996, Kamitani et al. 1998, Saitoh. . .

#### Figure Legends

Figure 1. Identification of the Shared Region of Overlap (S.R.O.) for hemizygous deletions in 21ql 1-q21 in NSCLCA Cytogenetic map, Not I long range physical. . .

the single key  
aminoacids in the Cys and His domains. Two reports show the localisations  
of the highly homologous sequences for the HAUSP gene to 3p21 (Kashuba, et al 1997) and 16p1 3 (Robinson, et al 1998), respectively.

A. 1992. Ubiquitin-specific proteases of *Saccharomyces cerevisiae*. J. Biol Chem 267:23364  
Baker, R.T., Wang, X-W., Woollatt, E., White, J.A. and Sutherlands, G.R. Identification, functional characterisation, and chromosomal localisation of USP15, a novel Human USP related to Unp Oncoprotein, and a systematic nomenclature for hUSP's. Genomics. . .

T., Saito, A... Suzuki, M., Shinomiya, H., Suzuki, T., Takahashi, E., Tanigami, A., Ichiyama, A., Chung, C.H., Nakamura, Y., and Tanaka, K. Identification and chromosomal assignment of USP1, a novel gene encoding a human ubiquitin-specific protease. Genomics, 54:155-158, 1998.

human chromosome 5q33-q34, UBH1, encodes a novel deubiquitinating enzyme. Genomics 49:411  
Haupt Y, Maya R, Kazaz A, Oren M (1 997) Mdm2 promotes the rapid degradation of p53, Nature 387:296  
Ichikawa, H., Hosoda, F., Arai, Y., Shimizu, K., Ohira, M., and Ohki, M. A. . . .

Sumegi J, Klein G, Zabarovsky ER, Kisselev L. 1997. Not1 linking/jumping clones of human chromosome 3: mapping of the TFRC, RA137 and HAUSP genes to regions rearranged in leukemia and deleted in solid tumours. FEBS Lett 419:181-185.

Assignment of herpesvirus-associated ubiquitin-specific protease gene HAUSP to human chromosome band 16p 13.3 by in situ hybridisation, Cytogenet. Cell Genet. 83:100.

=> file reg

COST IN U.S. DOLLARS

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FULL ESTIMATED COST

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=> E "HAUSP"/CN 25

|     |       |   |
|-----|-------|---|
| E1  | 1     | HAUSMANNITE, MAGNESIAN (MN2(MN0.5-0.9MG0.1-0.5)O4)/CN             |
| E2  | 1     | HAUSMANNITE, ZINCIAN (MN2(MN0.5-0.9ZN0.1-0.5)O4)/CN               |
| E3  | 0 --> | HAUSP/CN  |
| E4  | 1     | HAUSP PROTEASE/CN   |
| E5  | 1     | HAUSTELLUM SPECIFIC PROTEIN B (SARCOPHAGA PEREGRINA GENE HSPB)/CN |
| E6  | 1     | HAUTHANE HD 2334/CN   |
| E7  | 1     | HAUTHANE HD 2334, POLYMER WITH MONDUR TD 80 AND POLY-G 83-34/CN   |
| E8  | 1     | HAUTHANE HD 2001/CN   |
| E9  | 1     | HAUTHANE HD 4664/CN   |
| E10 | 1     | HAUTHANE L 2020/CN  |
| E11 | 1     | HAUTHAWAY IDA/CN  |
| E12 | 1     | HAUTOFOAM ES-AL/CN  |
| E13 | 1     | HAUTOFOAM ES-SI/CN  |
| E14 | 1     | HAUTOFOAM ES-TI/CN  |
| E15 | 1     | HAUTOFOAM ITO/CN  |
| E16 | 1     | HAUTOFOAM MS-Y/CN   |
| E17 | 1     | HAUTOFOAM NI/CN   |
| E18 | 1     | HAUTOFOAM SN/CN   |
| E19 | 1     | HAUTRIWAIC ACID/CN  |
| E20 | 1     | HAUTRIWAIC ACID $\Gamma$ -LACTONE/CN                              |
| E21 | 1     | HAUTRIWAIC ACID ACETATE/CN  |
| E22 | 1     | HAUTRIWAIC ACID LACTONE/CN  |
| E23 | 1     | HAUTRIWAIC ACID METHYL ESTER/CN                                   |
| E24 | 1     | HAUTRIWAIC ACID METHYL ESTER ACETATE/CN                           |
| E25 | 2     | HAUYNE/CN   |

=> S E4

L12 1 "HAUSP PROTEASE"/CN

=> DIS L12 1 SQIDE

THE ESTIMATED COST FOR THIS REQUEST IS 6.36 U.S. DOLLARS  
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L12 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2006 ACS on STN  
RN 109136-49-4 REGISTRY  
CN Proteinase, ubiquitin conjugate (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN DEN1 protease  
CN Deneddylase  
CN Deubiquination enzyme UBPl

CN Deubiquitinase  
 CN Deubiquitinating enzyme  
 CN Deubiquitinating enzyme DUB-2  
 CN HAUSP protease  
 CN ISG15-specific protease UBP43  
 CN Otubain 1  
 CN Polyubiquitin proteinase  
 CN Protease USP21  
 CN Proteinase, ubiquitin-fusion protein  
 CN Ubiquitin conjugate protease  
 CN Ubiquitin conjugate proteinase  
 CN Ubiquitin protease  
 CN Ubiquitin proteinase  
 CN Ubiquitin-fusion protein proteinase  
 CN Ubiquitin-specific processing protease  
 CN Ubiquitin-specific protease  
 CN Ubiquitin-specific protease 21  
 CN Ubiquitin-specific proteinase  
 CN UBP1 protease  
 DR 123175-78-0, 123175-79-1  
 MF Unspecified  
 CI MAN  
 SR CA  
 LC STN Files: AGRICOLA, BIOSIS, CA, CAPLUS, CIN, PROMT, TOXCENTER, USPAT2,  
 USPATFULL  
 DT.CA CAplus document type: Conference; Dissertation; Journal; Patent  
 RL.P Roles from patents: ANST (Analytical study); BIOL (Biological study);  
 MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC  
 (Process); PRP (Properties); USES (Uses)  
 RLD.P Roles for non-specific derivatives from patents: ANST (Analytical  
 study); BIOL (Biological study); PREP (Preparation); PRP (Properties);  
 USES (Uses)  
 RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological  
 study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP  
 (Properties); USES (Uses)  
 RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological  
 study); PROC (Process); PRP (Properties)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

610 REFERENCES IN FILE CA (1907 TO DATE)

7 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

617 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> E "USP7"/CN 25

E1 1 USP6NL PROTEIN (HUMAN CLONE IMAGE:4838780)/CN  
 E2 1 USP6NL PROTEIN (HUMAN CLONE MGC:41831 IMAGE:5296060)/CN  
 E3 0 --> USP7/CN  
 E4 1 USP8 PROTEIN (HUMAN CLONE IMAGE:6429817 GENE USP8)/CN  
 E5 1 USP8 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:5041516 GENE  
 USP8)/CN  
 E6 1 USP8-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:53905  
 IMAGE:5543601)/CN  
 E7 1 USP9X PROTEIN (HUMAN CLONE IMAGE:4538919 GENE USP9X)/CN  
 E8 1 USP9X PROTEIN (HUMAN CLONE IMAGE:6175281 GENE USP9X)/CN  
 E9 1 USPA (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE  
 USPA)/CN  
 E10 1 USPA (NITROBACTER WINOGRADSKYI STRAIN NB-255)/CN  
 E11 1 USPA (PASTEURELLA MULTOCIDA STRAIN IL1403 CLONE PM70 GENE  
 USPA)/CN  
 E12 3 USPA (PSEUDOMONAS SYRINGAE SYRINGAE STRAIN B728A)/CN  
 E13 1 USPA FAMILY PROTEIN (BURKHOLDERIA MALLEI STRAIN ATCC 23344)/CN  
 E14 1 USPA FAMILY PROTEIN (BURKHOLDERIA THAILANDENSIS STRAIN E264)/CN  
 E15 3 USPA PROTEIN (MANNHEIMIA SUCCINICIPRODUCENS STRAIN MBEL55E GENE  
 USPA)/CN

|     |   |  |
|-----|---|--|
| E16 | 1 | USPA-RELATED NUCLEOTIDE-BINDING PROTEIN (IDIOMARINA LOIHIENSIS STRAIN L2TR GENE USPA)/CN |
| E17 | 1 | USPA-RELATED NUCLEOTIDE-BINDING PROTEIN (IDIOMARINA LOIHIENSIS STRAIN L2TR)/CN           |
| E18 | 1 | USPALLATINE/CN   |
| E19 | 1 | USPALLATINE 6-ACETATE/CN   |
| E20 | 1 | USPALLATINE ACETATE/CN   |
| E21 | 1 | USPALLATINECINE/CN   |
| E22 | 1 | USPC (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE USPC)/CN                     |
| E23 | 1 | USPCA/CN   |
| E24 | 1 | USPE (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE USPE)/CN                     |
| E25 | 1 | USPIO/CN   |

=> E "USP 7"/CN 25

|     |       |  |
|-----|-------|--|
| E1  | 1     | USP 400/CN   |
| E2  | 1     | USP 400P/CN  |
| E3  | 0 --> | USP 7/CN   |
| E4  | 1     | USP 711/CN   |
| E5  | 1     | USP 90MD/CN  |
| E6  | 1     | USP DOMAIN (BACILLUS LICHENIFORMIS STRAIN ATCC 14580 GENE YXIE)/CN                       |
| E7  | 1     | USP10-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:53812 IMAGE:5542105)/CN                     |
| E8  | 1     | USP10-PROV PROTEIN (XENOPUS TROPICALIS CLONE MGC:89480 IMAGE:6991947 GENE USP10-PROV)/CN |
| E9  | 1     | USP11 PROTEIN (HUMAN CLONE IMAGE:2961383 GENE USP11)/CN                                  |
| E10 | 1     | USP11 PROTEIN (HUMAN CLONE IMAGE:4180680 GENE USP11)/CN                                  |
| E11 | 1     | USP12 PROTEIN (MOUSE CLONE MGC:55011 IMAGE:4500832)/CN                                   |
| E12 | 1     | USP12 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:5100708 GENE USP12)/CN                     |
| E13 | 1     | USP14 PROTEIN (MOUSE STRAIN 129/SV X 129/SV-CP CLONE MGC:60479 IMAGE:30006882)/CN        |
| E14 | 1     | USP15 PROTEIN (HUMAN CLONE IMAGE:4689787 GENE USP15)/CN                                  |
| E15 | 1     | USP15 PROTEIN (HUMAN CLONE IMAGE:4689787)/CN   |
| E16 | 1     | USP16 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:3586657 GENE USP16)/CN                  |
| E17 | 1     | USP16 PROTEIN (XENOPUS TROPICALIS CLONE IMAGE:6990711 GENE USP16)/CN                     |
| E18 | 1     | USP19 PROTEIN (HUMAN CLONE IMAGE:4478169 GENE USP19)/CN                                  |
| E19 | 1     | USP19 PROTEIN (HUMAN CLONE IMAGE:4859467 GENE USP19)/CN                                  |
| E20 | 1     | USP2 PROTEIN (HUMAN CLONE MGC:43844 IMAGE:5273400 GENE USP2)/CN                          |
| E21 | 1     | USP2 PROTEIN (HUMAN CLONE MGC:43844 IMAGE:5273400)/CN                                    |
| E22 | 1     | USP2 PROTEIN (MOUSE CLONE MGC:27630 IMAGE:4506362)/CN                                    |
| E23 | 1     | USP21 PROTEIN (HUMAN CLONE IMAGE:2958151 GENE USP21)/CN                                  |
| E24 | 1     | USP21 PROTEIN (HUMAN CLONE MGC:3394 IMAGE:2958151)/CN                                    |
| E25 | 1     | USP22 PROTEIN (MOUSE STRAIN C57BL/6 CLONE IMAGE:5686812 GENE USP22)/CN                   |

=> E "USP-7"/CN 25

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|----|-------|--|
| E1 | 1     | USP 90MD/CN  |
| E2 | 1     | USP DOMAIN (BACILLUS LICHENIFORMIS STRAIN ATCC 14580 GENE YXIE)/CN                       |
| E3 | 0 --> | USP-7/CN   |
| E4 | 1     | USP10-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:53812 IMAGE:5542105)/CN                     |
| E5 | 1     | USP10-PROV PROTEIN (XENOPUS TROPICALIS CLONE MGC:89480 IMAGE:6991947 GENE USP10-PROV)/CN |
| E6 | 1     | USP11 PROTEIN (HUMAN CLONE IMAGE:2961383 GENE USP11)/CN                                  |
| E7 | 1     | USP11 PROTEIN (HUMAN CLONE IMAGE:4180680 GENE USP11)/CN                                  |
| E8 | 1     | USP12 PROTEIN (MOUSE CLONE MGC:55011 IMAGE:4500832)/CN                                   |
| E9 | 1     | USP12 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:5100708 GENE USP12)/CN                     |



|     |   |  |
|-----|---|--|
| E10 | 1 | USP14 PROTEIN (MOUSE STRAIN 129/SV X 129/SV-CP CLONE MGC:60479<br>IMAGE:30006882)/CN |
| E11 | 1 | USP15 PROTEIN (HUMAN CLONE IMAGE:4689787 GENE USP15)/CN                              |
| E12 | 1 | USP15 PROTEIN (HUMAN CLONE IMAGE:4689787)/CN   |
| E13 | 1 | USP16 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:3586657 GENE<br>USP16)/CN           |
| E14 | 1 | USP16 PROTEIN (XENOPUS TROPICALIS CLONE IMAGE:6990711 GENE<br>USP16)/CN              |
| E15 | 1 | USP19 PROTEIN (HUMAN CLONE IMAGE:4478169 GENE USP19)/CN                              |
| E16 | 1 | USP19 PROTEIN (HUMAN CLONE IMAGE:4859467 GENE USP19)/CN                              |
| E17 | 1 | USP2 PROTEIN (HUMAN CLONE MGC:43844 IMAGE:5273400 GENE USP2)/CN                      |
| E18 | 1 | USP2 PROTEIN (HUMAN CLONE MGC:43844 IMAGE:5273400)/CN                                |
| E19 | 1 | USP2 PROTEIN (MOUSE CLONE MGC:27630 IMAGE:4506362)/CN                                |
| E20 | 1 | USP21 PROTEIN (HUMAN CLONE IMAGE:2958151 GENE USP21)/CN                              |
| E21 | 1 | USP21 PROTEIN (HUMAN CLONE MGC:3394 IMAGE:2958151)/CN                                |
| E22 | 1 | USP22 PROTEIN (MOUSE STRAIN C57BL/6 CLONE IMAGE:5686812 GENE<br>USP22)/CN            |
| E23 | 1 | USP24 PROTEIN (HUMAN CLONE IMAGE:4995223 GENE USP24)/CN                              |
| E24 | 1 | USP24 PROTEIN (HUMAN CLONE MGC:29848 IMAGE:4995223)/CN                               |
| E25 | 1 | USP24 PROTEIN (MOUSE STRAIN C57BL/6J CLONE MGC:41753<br>IMAGE:1381175)/CN            |

=> E 25

|     |   |   |
|-----|---|---|
| E26 | 1 | USP25 PROTEIN (HUMAN CLONE IMAGE:3920963)/CN  |
| E27 | 1 | USP25-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:81666 IMAGE:6864548<br>GENE USP25-PROV)/CN   |
| E28 | 1 | USP28 PROTEIN (HUMAN CLONE MGC:70638 IMAGE:6140725)/CN                                    |
| E29 | 1 | USP28 PROTEIN (MOUSE STRAIN C57BL/6 CLONE MGC:91310<br>IMAGE:6835629)/CN                  |
| E30 | 1 | USP3-PROV PROTEIN (XENOPUS TROPICALIS CLONE MGC:89592<br>IMAGE:6994758 GENE USP3-PROV)/CN |
| E31 | 1 | USP30 PROTEIN (HUMAN CLONE IMAGE:3536512 GENE USP30)/CN                                   |
| E32 | 1 | USP30 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:4911968 GENE<br>USP30)/CN                   |
| E33 | 1 | USP31 PROTEIN (HUMAN CLONE IMAGE:4650466)/CN  |
| E34 | 1 | USP32 PROTEIN (HUMAN CLONE IMAGE:6164242 GENE USP32)/CN                                   |
| E35 | 1 | USP32 PROTEIN (HUMAN CLONE IMAGE:6502630)/CN  |
| E36 | 1 | USP32 PROTEIN (MOUSE CLONE IMAGE:5369173 GENE USP32)/CN                                   |
| E37 | 1 | USP32 PROTEIN (MOUSE STRAIN C57BL/6 CLONE IMAGE:5706776 GENE<br>USP32)/CN                 |
| E38 | 1 | USP33 PROTEIN (MOUSE CLONE IMAGE:5359218 GENE USP33)/CN                                   |
| E39 | 1 | USP33 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:5066137 GENE<br>USP33)/CN                |
| E40 | 1 | USP33 PROTEIN (MOUSE STRAIN MIX FVB/N, C57BL/6J CLONE<br>IMAGE:3491447 GENE USP33)/CN     |
| E41 | 1 | USP34 PROTEIN (HUMAN CLONE IMAGE:3846921 GENE USP34)/CN                                   |
| E42 | 1 | USP34 PROTEIN (HUMAN CLONE IMAGE:5426830)/CN  |
| E43 | 1 | USP34 PROTEIN (MOUSE STRAIN C57BL/6 CLONE IMAGE:5708348 GENE<br>USP34)/CN                 |
| E44 | 1 | USP34 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:3983228 GENE<br>USP34)/CN                |
| E45 | 1 | USP36 PROTEIN (HUMAN CLONE MGC:87250 IMAGE:30345255)/CN                                   |
| E46 | 1 | USP39 PROTEIN (HUMAN CLONE IMAGE:3050936 GENE USP39)/CN                                   |
| E47 | 1 | USP4 PROTEIN (MOUSE STRAIN C57BL/6J CLONE IMAGE:30433369 GENE<br>USP4)/CN                 |
| E48 | 1 | USP4 PROTEIN (MOUSE STRAIN C57BL/6J CLONE IMAGE:30457732 GENE<br>USP4)/CN                 |
| E49 | 1 | USP4 PROTEIN (MOUSE STRAIN CZECH II CLONE MGC:18374<br>IMAGE:3661165)/CN                  |
| E50 | 1 | USP42 PROTEIN (HUMAN CLONE IMAGE:30343487 GENE USP42)/CN                                  |

=> E 25

|     |   |   |
|-----|---|---|
| E51 | 1 | USP43 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:3711168 GENE<br>USP43)/CN |
|-----|---|---|

|                           |   |   |
|---------------------------|---|---|
| E52                       | 1 | USP43 PROTEIN (MOUSE STRAIN FVB/N CLONE MGC:38313               |
| IMAGE:5343429)/CN         |   |   |
| E53                       | 1 | USP44 PROTEIN (HUMAN CLONE MGC:26981 IMAGE:4825887)/CN          |
| E54                       | 1 | USP45 PROTEIN (HUMAN CLONE MGC:14793 IMAGE:4047601)/CN          |
| E55                       | 1 | USP46-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:53480              |
| IMAGE:5572291)/CN         |   |   |
| E56                       | 1 | USP47 PROTEIN (HUMAN CLONE IMAGE:3350895 GENE USP47)/CN         |
| E57                       | 1 | USP47 PROTEIN (HUMAN CLONE IMAGE:5575723)/CN                    |
| E58                       | 1 | USP47 PROTEIN (MOUSE STRAIN C57BL/6 CLONE IMAGE:6813792)/CN     |
| E59                       | 1 | USP47 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:5068518 GENE   |
| USP47)/CN                 |   |   |
| E60                       | 1 | USP47 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:4211195 GENE      |
| USP47)/CN                 |   |   |
| E61                       | 1 | USP48 PROTEIN (HUMAN CLONE IMAGE:3897418 GENE USP48)/CN         |
| E62                       | 1 | USP48 PROTEIN (HUMAN CLONE IMAGE:4650466 GENE USP48)/CN         |
| E63                       | 1 | USP48 PROTEIN (HUMAN CLONE IMAGE:4650466)/CN                    |
| E64                       | 1 | USP48 PROTEIN (HUMAN CLONE IMAGE:6514749 GENE USP48)/CN         |
| E65                       | 1 | USP48 PROTEIN (MOUSE STRAIN FVB/N CLONE MGC:25724 IMAGE:3979404 |
| GENE USP48)/CN            |   |   |
| E66                       | 1 | USP49 PROTEIN (MOUSE STRAIN C57BL/6 CLONE MGC:63299             |
| IMAGE:6408678)/CN         |   |   |
| E67                       | 1 | USP5 PROTEIN (HUMAN CLONE MGC:1586 IMAGE:3506801)/CN            |
| E68                       | 1 | USP51 PROTEIN (HUMAN CLONE IMAGE:5391875 GENE USP51)/CN         |
| E69                       | 1 | USP52 PROTEIN (MOUSE STRAIN CZECH II CLONE IMAGE:3591885 GENE   |
| USP52)/CN                 |   |   |
| E70                       | 1 | USP52 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:4988880)/CN       |
| E71                       | 1 | USP53 PROTEIN (HUMAN CLONE MGC:22206 IMAGE:4082351)/CN          |
| E72                       | 1 | USP53 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:4236151 GENE      |
| USP53)/CN                 |   |   |
| E73                       | 1 | USP54 PROTEIN (HUMAN CLONE IMAGE:6148876 GENE USP54)/CN         |
| E74                       | 1 | USP54 PROTEIN (MOUSE STRAIN CD1 CLONE IMAGE:30088889 GENE       |
| USP54)/CN                 |   |   |
| E75                       | 1 | USP54 PROTEIN (MOUSE STRAIN CD1 CLONE IMAGE:30096132 GENE       |
| USP54)/CN                 |   |   |
| => E 25                   |   |   |
| E76                       | 1 | USP6 N-TERMINAL LIKE (MOUSE CLONE MGC:57018 IMAGE:6466270 GENE  |
| USP6NL)/CN                |   |   |
| E77                       | 1 | USP6NL PROTEIN (HUMAN CLONE IMAGE:4047207 GENE USP6NL)/CN       |
| E78                       | 1 | USP6NL PROTEIN (HUMAN CLONE IMAGE:4838780)/CN                   |
| E79                       | 1 | USP6NL PROTEIN (HUMAN CLONE MGC:41831 IMAGE:5296060)/CN         |
| E80                       | 1 | USP8 PROTEIN (HUMAN CLONE IMAGE:6429817 GENE USP8)/CN           |
| E81                       | 1 | USP8 PROTEIN (MOUSE STRAIN FVB/N CLONE IMAGE:5041516 GENE       |
| USP8)/CN                  |   |   |
| E82                       | 1 | USP8-PROV PROTEIN (XENOPUS LAEVIS CLONE MGC:53905               |
| IMAGE:5543601)/CN         |   |   |
| E83                       | 1 | USP9X PROTEIN (HUMAN CLONE IMAGE:4538919 GENE USP9X)/CN         |
| E84                       | 1 | USP9X PROTEIN (HUMAN CLONE IMAGE:6175281 GENE USP9X)/CN         |
| E85                       | 1 | USPA (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE     |
| USPA)/CN                  |   |   |
| E86                       | 1 | USPA (NITROBACTER WINOGRADSKYI STRAIN NB-255)/CN                |
| E87                       | 1 | USPA (PASTEURELLA MULTOCIDA STRAIN IL1403 CLONE PM70 GENE       |
| USPA)/CN                  |   |   |
| E88                       | 3 | USPA (PSEUDOMONAS SYRINGAE SYRINGAE STRAIN B728A)/CN            |
| E89                       | 1 | USPA FAMILY PROTEIN (BURKHOLDERIA MALLEI STRAIN ATCC 23344)/CN  |
| E90                       | 1 | USPA FAMILY PROTEIN (BURKHOLDERIA THAILANDENSIS STRAIN E264)/CN |
| E91                       | 3 | USPA PROTEIN (MANNHEIMIA SUCCINICIPRODUCENS STRAIN MBEL55E GENE |
| USPA)/CN                  |   |   |
| E92                       | 1 | USPA-RELATED NUCLEOTIDE-BINDING PROTEIN (IDIOMARINA LOIHIENSIS  |
| STRAIN L2TR GENE USPA)/CN |   |   |
| E93                       | 1 | USPA-RELATED NUCLEOTIDE-BINDING PROTEIN (IDIOMARINA LOIHIENSIS  |
| STRAIN L2TR)/CN           |   |   |
| E94                       | 1 | USPALLATINE/CN  |
| E95                       | 1 | USPALLATINE 6-ACETATE/CN  |

|      |   |   |
|------|---|---|
| E96  | 1 | USPALLATINE ACETATE/CN  |
| E97  | 1 | USPALLATINECINE/CN  |
| E98  | 1 | USPC (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE<br>USPC)/CN |
| E99  | 1 | USPCA/CN  |
| E100 | 1 | USPE (MYCOBACTERIUM AVIUM PARATUBERCULOSIS STRAIN K-10 GENE<br>USPE)/CN |

=> file medline

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

8.42

54.46

FILE 'MEDLINE' ENTERED AT 15:19:32 ON 20 SEP 2006

FILE LAST UPDATED: 19 Sep 2006 (20060919/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>).

See also:

<http://www.nlm.nih.gov/mesh/>

[http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\\_mesh.html](http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html)

[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_med\\_data\\_changes.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html)

[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_2006\\_MeSH.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html)

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s HAUSP or (USP ( ) 7)

39 HAUSP

4983 USP

37 USPS

5006 USP

(USP OR USPS)

1527014 7

0 USP (W) 7

L13 39 HAUSP OR (USP (W) 7)

=> s HAUSP or (USP7)

39 HAUSP

47 USP7

L14 55 HAUSP OR (USP7)

=> s MDM2

L15 2699 MDM2

=> s l15 and l14

L16 18 L15 AND L14

=> s l16 not py>2002

2271354 PY>2002

(PY>20029999)

L17 1 L16 NOT PY>2002

=> d ibib

L17 ANSWER 1 OF 1 MEDLINE on STN  
 ACCESSION NUMBER: 2002212418 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 11923872  
 TITLE: Deubiquitination of p53 by HAUSP is an important pathway for p53 stabilization.  
 AUTHOR: Li Muyang; Chen Delin; Shiloh Ariel; Luo Jianyuan; Nikolaev Anatoly Y; Qin Jun; Gu Wei  
 CORPORATE SOURCE: Institute for Cancer Genetics, and Department of Pathology, College of Physicians & Surgeons, Columbia University, 1150 St Nicholas Avenue, New York, New York 10032, USA.  
 SOURCE: Nature, (2002 Apr 11) Vol. 416, No. 6881, pp. 648-53.  
 Electronic Publication: 2002-03-31.  
 Journal code: 0410462. ISSN: 0028-0836.  
 PUB. COUNTRY: England; United Kingdom  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 200205  
 ENTRY DATE: Entered STN: 12 Apr 2002  
 Last Updated on STN: 18 May 2002  
 Entered Medline: 17 May 2002

=> d abs

L17 ANSWER 1 OF 1 MEDLINE on STN  
 AB The p53 tumour suppressor is a short-lived protein that is maintained at low levels in normal cells by Mdm2-mediated ubiquitination and subsequent proteolysis. Stabilization of p53 is crucial for its tumour suppressor function. However, the precise mechanism by which ubiquitinated p53 levels are regulated in vivo is not completely understood. By mass spectrometry of affinity-purified p53-associated factors, we have identified herpesvirus-associated ubiquitin-specific protease (HAUSP) as a novel p53-interacting protein. HAUSP strongly stabilizes p53 even in the presence of excess Mdm2, and also induces p53-dependent cell growth repression and apoptosis. Significantly, HAUSP has an intrinsic enzymatic activity that specifically deubiquitinates p53 both in vitro and in vivo. In contrast, expression of a catalytically inactive point mutant of HAUSP in cells increases the levels of p53 ubiquitination and destabilizes p53. These findings reveal an important mechanism by which p53 can be stabilized by direct deubiquitination and also imply that HAUSP might function as a tumour suppressor in vivo through the stabilization of p53.

=> d his

(FILE 'HOME' ENTERED AT 15:06:58 ON 20 SEP 2006)

FILE 'CAPLUS' ENTERED AT 15:07:09 ON 20 SEP 2006

L1 21 S HAUSP AND MDM2  
 L2 6 S L1 NOT PY>2004  
 L3 40 S USP7  
 L4 8 S L3 AND MDM2

FILE 'PCTFULL' ENTERED AT 15:12:56 ON 20 SEP 2006

L5 37 S USP7  
 L6 34 S HAUSP  
 L7 59 S L6 OR L5  
 L8 18 S MDM2 AND L7  
 L9 532010 S SCREEN? OR IDENT?  
 L10 18 S L9 AND L8  
 L11 5 S L10 NOT PY>2002

FILE 'REGISTRY' ENTERED AT 15:17:14 ON 20 SEP 2006

L12           E "HAUSP"/CN 25  
              1 S E4  
              E "USP7"/CN 25  
              E "USP 7"/CN 25  
              E "USP-7"/CN 25

FILE 'MEDLINE' ENTERED AT 15:19:32 ON 20 SEP 2006

L13           39 S HAUSP OR (USP () 7)  
L14           55 S HAUSP OR (USP7)  
L15           2699 S MDM2  
L16           18 S L15 AND L14  
L17           1 S L16 NOT PY>2002

=> file pctfull

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

3.73

58.19

FILE 'PCTFULL' ENTERED AT 15:24:49 ON 20 SEP 2006

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FILE LAST UPDATED:           18 SEP 2006           <20060918/UP>  
MOST RECENT UPDATE WEEK:       200637            <200637/EW>  
FILE COVERS 1978 TO DATE

>>> IMAGES ARE AVAILABLE ONLINE AND FOR EMAIL-PRINTS <<<

>>> NEW IPC8 DATA AND FUNCTIONALITY NOW AVAILABLE IN THIS FILE.

SEE

<http://www.stn-international.de/stndatabases/details/ipc-reform.html> >>>

>>> FOR CHANGES IN PCTFULL PLEASE SEE HELP CHANGE  
(last updated April 10, 2006) <<<

>>> NEW PRICES IN PCTFULL AS OF 01 JULY 2006. FOR DETAILS,  
PLEASE SEE HELP COST <<<

=> d l11 ibib

L11    ANSWER 1 OF 5           PCTFULL    COPYRIGHT 2006 Univentio on STN  
ACCESSION NUMBER:           2002070742 PCTFULL   ED 20020926   EW 200237  
TITLE (ENGLISH):           METHOD FOR THE DEVELOPMENT OF GENE PANELS FOR  
                              DIAGNOSTIC AND THERAPEUTIC PURPOSES BASED ON THE  
                              EXPRESSION AND METHYLATION STATUS OF THE GENES  
TITLE (FRENCH):            PROCEDE DE MISE AU POINT DE GROUPE D'ECHANTILLONS DE  
                              GENES A DES FINS DE DIAGNOSTIC ET DE THERAPIE QUI SONT  
                              BASEES SUR L'EXPRESSION ET L'ETAT DE METHYLATION DES  
                              GENES  
INVENTOR(S):               OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
                              DE;  
                              BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahndorf, DE  
PATENT ASSIGNEE(S):        EPIGENOMICS AG, Kastanienallee 24, 10435 Berlin, DE [DE,  
                              DE]  
AGENT:                    SCHOHE, Stefan\$, Boehmert & Boehmert, Pettenkoferstr.  
                              20-22, 80336 Muenchen\$, DE  
LANGUAGE OF FILING:        English  
LANGUAGE OF PUBL.:         English  
DOCUMENT TYPE:             Patent  
PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002070742 | A1   | 20020912 |

## DESIGNATED STATES

W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
 CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
 SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW

RW (ARIPO):

GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW

RW (EAPO):

AM AZ BY KG KZ MD RU TJ TM

RW (EPO):

AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR

RW (OAPI):

BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG

APPLICATION INFO.:

WO 2002-EP2255 A 20020301

PRIORITY INFO.:

US 2001-60/272,549 20010301

=&gt; d 111 ibib 1-5

L11 ANSWER 1 OF 5

ACCESSION NUMBER:

PCTFULL COPYRIGHT 2006 Univentio on STN

TITLE (ENGLISH):

2002070742 PCTFULL ED 20020926 EW 200237

TITLE (FRENCH):

METHOD FOR THE DEVELOPMENT OF GENE PANELS FOR  
 DIAGNOSTIC AND THERAPEUTIC PURPOSES BASED ON THE  
 EXPRESSION AND METHYLATION STATUS OF THE GENES  
 PROCEDE DE MISE AU POINT DE GROUPES D'ECHANTILLONS DE  
 GENES A DES FINS DE DIAGNOSTIC ET DE THERAPIE QUI SONT  
 BASES SUR L'EXPRESSION ET L'ETAT DE METHYLATION DES  
 GENES

INVENTOR(S):

OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
 DE;

PATENT ASSIGNEE(S):

BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahndorf, DE  
 EPIGENOMICS AG, Kastanienalle 24, 10435 Berlin, DE [DE,  
 DE]

AGENT:

SCHOHE, Stefan\$, Boehmert & Boehmert, Pettenkoferstr.  
 20-22, 80336 Muenchen\$, DE

LANGUAGE OF FILING:

English

LANGUAGE OF PUBL.:

English

DOCUMENT TYPE:

Patent

PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| -----         |      |          |
| WO 2002070742 | A1   | 20020912 |

## DESIGNATED STATES

W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
 CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
 SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW

RW (ARIPO):

GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW

RW (EAPO):

AM AZ BY KG KZ MD RU TJ TM

RW (EPO):

AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR

RW (OAPI):

BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG

APPLICATION INFO.:

WO 2002-EP2255 A 20020301

PRIORITY INFO.:

US 2001-60/272,549 20010301

L11 ANSWER 2 OF 5

ACCESSION NUMBER:

PCTFULL COPYRIGHT 2006 Univentio on STN

TITLE (ENGLISH):

2002070741 PCTFULL ED 20020926 EW 200237

TITLE (FRENCH):

METHODS, SYSTEMS AND COMPUTER PROGRAM PRODUCTS FOR  
 DETERMINING THE BIOLOGICAL EFFECT AND/OR ACTIVITY OF  
 DRUGS, CHEMICAL SUBSTANCES AND/OR PHARMACEUTICAL  
 COMPOSITIONS BASED ON THEIR EFFECT ON THE METHYLATION  
 STATUS OF THE DNA  
 PROCEDES, SYSTEMES ET PRODUITS PROGRAMMES INFORMATIQUES  
 PERMETTANT DE DETERMINER L'EFFET BIOLOGIQUE ET/OU  
 L'ACTIVITE DE MEDICAMENTS, DE SUBSTANCES CHIMIQUES

INVENTOR(S): ET/OU DE COMPOSITIONS PHARMACEUTIQUES, SUR LA BASE DE  
LEUR EFFET SUR L'ETAT DE METHYLATION DE L'ADN  
OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
DE;  
PATENT ASSIGNEE(S): BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahnsdorf, DE  
EPIGENOMICS AG, Kastanienallee 24, 10435 Berlin, DE  
[DE, DE]  
AGENT: SCHOHE, Stefan\$, Boehmert & Boehmert,  
Pettenkoferstrasse 20-22, 80336 Muenchen\$, DE  
LANGUAGE OF FILING: English  
LANGUAGE OF PUBL.: English  
DOCUMENT TYPE: Patent  
PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002070741 | A2   | 20020912 |

DESIGNATED STATES  
W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW  
GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW  
AM AZ BY KG KZ MD RU TJ TM  
AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
TR

RW (ARIPO):  
RW (EAPO):  
RW (EPO):  
RW (OAPI):  
APPLICATION INFO.:  
PRIORITY INFO.:

BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG  
WO 2002-EP2254 A 20020301  
US 2001-60/272,484 20010301

L11 ANSWER 3 OF 5  
ACCESSION NUMBER:  
TITLE (ENGLISH):  
TITLE (FRENCH):  
INVENTOR(S):

PCTFULL COPYRIGHT 2006 Univentio on STN  
2002057414 PCTFULL ED 20020801 EW 200230  
LEUKOCYTE EXPRESSION PROFILING  
EVALUATION DU NIVEAU D'EXPRESSION LEUCOCYTAIRE  
WOHLGEMUTH, Jay, 664 Hamilton Avenue, Palo Alto, CA  
94301, US [US, US];  
FRY, Kirk, 2604 Ross Road, Palo Alto, CA 94303, US [US,  
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MATCUK, George, 141C Escondido Village, Stanford, CA  
94305, US [US, US];  
ALTMAN, Peter, 717 Evelyn Avenue, Albany, CA 94706, US  
[US, US];  
PRENTICE, James, 120 Dolores Street, San Francisco, CA  
94103, US [US, US];  
PHILLIPS, Julie, 1090 Mirador Terrace, Pacifica, CA  
94044, US [US, US];  
LY, Ngoc, 2000 Crystal Springs Road 15-14, San Bruno,  
CA 94066, US [US, US];  
WOODWARD, Robert, 1828 Rheem Court, Pleasanton, CA  
94588, US [US, US];  
QUENTERMOUS, Thomas, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US];  
JOHNSON, Frances, 44 El Rey Road, Portola Valley, CA  
94028, US [US, US]  
PATENT ASSIGNEE(S): BIOCARDIA, INC., 384 Oyster Point Boulevard, #4, South  
San Francisco, CA 94080, US [US, US], for all  
designates States except US;  
WOHLGEMUTH, Jay, 664 Hamilton Avenue, Palo Alto, CA  
94301, US [US, US], for US only;  
FRY, Kirk, 2604 Ross Road, Palo Alto, CA 94303, US [US,  
US], for US only;  
MATCUK, George, 141C Escondido Village, Stanford, CA  
94305, US [US, US], for US only;  
ALTMAN, Peter, 717 Evelyn Avenue, Albany, CA 94706, US

[US, US], for US only;  
 PRENTICE, James, 120 Dolores Street, San Francisco, CA  
 94103, US [US, US], for US only;  
 PHILLIPS, Julie, 1090 Mirador Terrace, Pacifica, CA  
 94044, US [US, US], for US only;  
 LY, Ngoc, 2000 Crystal Springs Road 15-14, San Bruno,  
 CA 94066, US [US, US], for US only;  
 WOODWARD, Robert, 1828 Rheem Court, Pleasanton, CA  
 94588, US [US, US], for US only;  
 QUERTERMOUS, Thomas, 44 El Rey Road, Portola Valley, CA  
 94028, US [US, US], for US only;  
 JOHNSON, Frances, 44 El Rey Road, Portola Valley, CA  
 94028, US [US, US], for US only;  
 WARD, Michael, R.\$, Morrison & Foerster LLP, 425 Market  
 Street, San Francisco, CA 94105-2482\$, US

AGENT:

LANGUAGE OF FILING:  
 LANGUAGE OF PUBL.:  
 DOCUMENT TYPE:  
 PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002057414 | A2   | 20020725 |

DESIGNATED STATES

W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
 CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ PH PL PT RO RU SD SE SG SI SK  
 SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

RW (ARIPO):

GH GM KE LS MW MZ SD SL SZ TZ UG ZW

RW (EAPO):

AM AZ BY KG KZ MD RU TJ TM

RW (EPO):

AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR

RW (OAPI):

BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG

APPLICATION INFO.:

WO 2001-US47856 A 20011022

PRIORITY INFO.:

US 2000-60/241,994 20001020

US 2001-60/296,764 20010608

L11 ANSWER 4 OF 5

PCTFULL COPYRIGHT 2006 Univention on STN

ACCESSION NUMBER:

2000079267 PCTFULL ED 20020515

TITLE (ENGLISH):

TREATMENT OF CANCER

TITLE (FRENCH):

TRAITEMENT ANTICANCEREUX

INVENTOR(S):

NIZETIC, Dean;

PATENT ASSIGNEE(S):

GROET, JuergenRP : GILL JENNINGS & EVERY  
 SCHOOL OF PHARMACY, UNIVERSITY OF LONDON;  
 NIZETIC, Dean;  
 GROET, Juergen

LANGUAGE OF PUBL.:

English

DOCUMENT TYPE:

Patent

PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2000079267 | A2   | 20001228 |

DESIGNATED STATES

W:

AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU  
 CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN  
 IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK  
 MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM  
 TR TT TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD  
 SL SZ TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY  
 DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ CF CG  
 CI CM GA GN GW ML MR NE SN TD TG

APPLICATION INFO.:

WO 2000-GB2446 A 20000622

PRIORITY INFO.:

GB 2000-0008161.2 20000403

GB 1999-9914589.8 19990622



L11 ANSWER 5 OF 5 PCTFULL COPYRIGHT 2006 Univentio on STN  
 ACCESSION NUMBER: 2000073479 PCTFULL ED 20020515  
 TITLE (ENGLISH): A COMBINED GROWTH FACTOR-DELETED AND THYMIDINE  
 KINASE-DELETED VACCINIA VIRUS VECTOR  
 TITLE (FRENCH): VECTEUR DU VIRUS DE LA VACCINE COMBINANT DELETION DU  
 FACTEUR DE CROISSANCE ET DELETION DE THYMIDINE KINASE  
 INVENTOR(S): MCCART, J., Andrea;  
 BARTLETT, David, L.;  
 MOSS, BernardRP : NATAUPSKY, Steven, J.  
 PATENT ASSIGNEE(S): THE GOVERNMENT OF THE UNITED STATES OF AMERICA, as  
 represented by THE SECRETARY, DEPARTMENT OF HEALTH AND  
 HUMAN SERVICES;  
 MCCART, J., Andrea;  
 BARTLETT, David, L.;  
 MOSS, Bernard  
 LANGUAGE OF PUBL.: English  
 DOCUMENT TYPE: Patent  
 PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2000073479 | A1   | 20001207 |

DESIGNATED STATES  
 W:

AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ  
 DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS  
 JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN  
 MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT  
 TZ UA UG US UZ VN YU ZA ZW GH GM KE LS MW MZ SD SL SZ  
 TZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE CH CY DE DK  
 ES FI FR GB GR IE IT LU MC NL PT SE BF BJ CF CG CI CM  
 GA GN GW ML MR NE SN TD TG

APPLICATION INFO.: WO 2000-US14679 A 20000526  
 PRIORITY INFO.: US 1999-60/137,126 19990528

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L11 ANSWER 2 OF 5 PCTFULL COPYRIGHT 2006 Univentio on STN  
 ACCESSION NUMBER: 2002070741 PCTFULL ED 20020926 EW 200237  
 TITLE (ENGLISH): METHODS, SYSTEMS AND COMPUTER PROGRAM PRODUCTS FOR  
 DETERMINING THE BIOLOGICAL EFFECT AND/OR ACTIVITY OF  
 DRUGS, CHEMICAL SUBSTANCES AND/OR PHARMACEUTICAL  
 COMPOSITIONS BASED ON THEIR EFFECT ON THE METHYLATION  
 STATUS OF THE DNA  
 TITLE (FRENCH): PROCEDES, SYSTEMES ET PRODUITS PROGRAMMES INFORMATIQUES  
 PERMETTANT DE DETERMINER L'EFFET BIOLOGIQUE ET/OU  
 L'ACTIVITE DE MEDICAMENTS, DE SUBSTANCES CHIMIQUES  
 ET/OU DE COMPOSITIONS PHARMACEUTIQUES, SUR LA BASE DE  
 LEUR EFFET SUR L'ETAT DE METHYLATION DE L'ADN  
 INVENTOR(S): OLEK, Alexander, Schroederstrasse 13/2, 10115 Berlin,  
 DE;  
 PATENT ASSIGNEE(S): BERLIN, Kurt, Marienkaeferweg 4, 14532 Stahnsdorf, DE  
 EPIGENOMICS AG, Kastanienallee 24, 10435 Berlin, DE  
 [DE, DE]  
 AGENT: SCHOHE, Stefan\$, Boehmert & Boehmert,  
 Pettenkoferstrasse 20-22, 80336 Muenchen\$, DE  
 LANGUAGE OF FILING: English  
 LANGUAGE OF PUBL.: English  
 DOCUMENT TYPE: Patent  
 PATENT INFORMATION:

| NUMBER        | KIND | DATE     |
|---------------|------|----------|
| WO 2002070741 | A2   | 20020912 |

DESIGNATED STATES

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR  
 CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID  
 IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD  
 MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI  
 SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW  
 RW (ARIPO): GH GM KE LS MW MZ SD SL SZ TZ UG ZM ZW  
 RW (EAPO): AM AZ BY KG KZ MD RU TJ TM  
 RW (EPO): AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE  
 TR  
 RW (OAPI): BF BJ CF CG CI CM GA GN GQ GW ML MR NE SN TD TG  
 APPLICATION INFO.: WO 2002-EP2254 A 20020301  
 PRIORITY INFO.: US 2001-60/272,484 20010301

DETD . . . since in most of the cases an effective  
 drug/treatment has to be found very rapidly,  
 Furthermore, such developments currently involve very cost-intensive  
 screening procedures  
 until a particularly suited compound (often called Jead&quot;-cornpound)  
 is found which could  
 then serve as a chemical basis for an effective treatment.

of course, alternative  
 treatments for already known diseases. Furthermore, the need exists for  
 a reliable, cost-  
 effective, fast and autornateable method for screening such  
 new effective compounds.

2. Screening for new biologically active compounds using  
 ,combinatorial chemistry&quot;  
 The method of combinatorial chemistry is described as a profound change  
 in the. . . AT, et  
 al. ,Search and discovery strategies for biotechnology: the paradigm  
 shift.&quot; Microbiol Mol  
 Biol Rev 2000 Sep;64(3):573-606)  
 In general, combinatorial chemistry involves screening of a  
 specific (or a set of specific)  
 compound with a vast number of otential biological candidate substances  
 for example, pro-  
 p  
 teins) that might interact with the compound. Interacting partners are  
 selected and used for  
 further screening. Initially screened and isolated  
 compounds can be used as Jead&quot;-  
 compounds for the development of biologically active compounds useful  
 for treatment of dis-  
 eases.

potential utility for the treatment of  
 conditions involving cerebral hypoxia.&quot; Life Sci 2000 Aug 1  
 1;67(12):1389-96) describe the  
 use of HTS (high-throughput screening) libraries for  
 reevaluation of the pharmacologic prop-  
 erties of substances such as extract from the leaves of Ginkgo biloba  
 Linne (form.. . .

Although the method of combinatorial chemistry exhibits several  
 advantages in comparison to  
 conventional methods for screening for biologically effective  
 compounds which are useful for  
 the development of new medicaments, there are still several drawbacks  
 associated with this  
 method.

The screening of a combinatorial chemistry library involves a

screening for a multitude of different possible reactions and/or interactions of the compounds to be analysed with the interacting partners. Therefore, the reaction conditions are assumed crucial for the result of the

screening. In particular, a compound which shows an interaction with a target in such a combinatorial assay in vitro might exhibit. . . prediction of an effective compound very difficult and unreliable. As a result, an interaction in an in vitro combinatorial chemistry screening assay can always only give a hint for a potential biological function of the screened compound in vivo.

As a result, combinatorial chemistry screening involves a necessary second step; once a potential target/lead compound has been identified/found, the biological effect still has to be confirmed/determined in an in vivo context. This makes compound identification using this method unpredictable, slow and costly.

only individual regions up to approximately 3000 base pairs in length have been examined, and an overall examination of cells to identify thousands of possible methylation events is not possible. However, this method is not capable of reliably analyzing minute fragments from small. . .

Burkitt's lymphoma: molecular analysis of primary tumor tissue; Blood 1998 Feb 15;91(4):13 73-8 1)  
- Wilms tumor (Kleyanova EV et al. "Identification of a tumor-specific methylation site in the Wilms tumor suppressor gene"; Oncogene 1998 Feb 12;16(6):713-20)  
- Prader-Willi/Angelman syndrome (Zeschnigh et al. "Imprinted. . .

The present invention uses the modifications in the methylation pattern of the DNA for

screening of biologically effective substances. In general, the invention uses the fact that the biological effect of a potentially biologically effective drug, . . .

The invention has several advantages in comparison to other screening methods; in particular combinatorial chemistry. First, the reaction conditions of the drug, chemical substance or pharmaceutical composition with the biological test system. . .

Second, the analysis of the methylation pattern of the DNA allows screening of the in vivo effect of the substance in a one-step procedure using one controllable reaction (namely, the bisulfite treatment in order. . .

Thirdly, screening for potential lead-compounds becomes less time consuming and less costly, since the complete screening and analysis procedure can be automated.

Fourth, the inventive method allows the inclusion of personal data into the selection/analysis

procedure which allows for a personalised screening of drugs, chemical substances or pharmaceutical compositions.

In a further preferred method according to the invention, the biological samples A and B are obtained from the identical individual, tissue, cell or other biological material.

or  
pharmaceutical composition. This allows the use of the inventive method to monitor and/or modify an already employed treatment regimen and to screen for unwanted side effects of the initially employed drugs, chemical substances or pharmaceutical compositions which leads to a strictly ,personalised" medicament. . .

cytosine methylation sites is analysed in parallel. The analysis of a multitude of sites in parallel allows for both an effective screening and a statistically highly relevant result of the method.

one to directly connect the tested drug, chemical substance or pharmaceutical composition with an effect on those genes and therefore allow the identification of possibly valuable new lead compounds as well as therapeutically important compounds.

In one embodiment, the method of the invention is characterised in that the identical biological sample, different biological samples or a combination thereof is used in steps a) and/or b).

#### Example 2

##### Screening of a peptide library

A peptide library was prepared in a 96-well culture plate which contained overlapping peptide fragments derived from the. . .

micro arrays representing 256 CpG and the methylation statuses of the CpGs were analysed according to a method described in Example 3

##### Screening of a fractionated plant crude extract

In order to analyse the anti-metastatic effect of *Celosia argentea* seed extracts (CAE), which have traditionally. . .

(CD47 anti-gen (Rh-related antigen, integrin-associated signal transducer); CD48 (CD48 antigen (B-cell membrane protein); CD53 (CD53 antigen); CD59 (CD59 antigen p18-20 (antigen identified by monoclonal antibodies 16.3A5, EJ16, EBO, EL32 and G344); CD63 (CD63 antigen (melanoma I antigen); CD68 (CD68 antigen); CD7 (CD7 antigen. . . LAMA4 (Laminin, alpha 4); LAMA5 (Laminin, alpha 5); LY64 (Lymphocyte antigen 64 (mouse) homolog, radioprotective, 105kD); LYZ (Lysozyme (renal amyloidosis)); MDUI (Antigen identified by monoclonal antibodies 4F2, TRAI.10, TROP4, and T43); MET (Met proto-oncogene (hepatocyte growth factor

receptor)); MIC2 (Antigen identified by monoclonal antibodies 12E7, F21 and 013); MICA (MHC class I polypeptide-related sequence A); MME (Membrane metallo-endopeptidase (neutral endopeptidase, enkephalinase, . . .

I (BCL2-related)); MCM4 (Minichromosome maintenance deficient (S. cerevisiae) 4); MEKK3 (MAP/ERK kinase kinase 3); MEKK5 (MAP/ERK kinase kinase 5); MKI67 (Antigen identified by monoclonal antibody Ki-67); MSTIR (Macrophage stimulating 1 receptor (c-met-related tyrosine kinase)); NCK1 (NCK adaptor protein 1); NEK3 (NIMA (never. . .

of split); AFD I (Acrofacial dysostosis 1, Nager type); AGC I (Aggrecan I (chondroitin sulfate proteoglycan 1, large aggregating proteoglycan, antigen identified by monoclonal antibody AO 1 22)); AH02 (Albright hereditary osteodystrophy-2); A1113 (Amelogenesis imperfecta 3, hypoplasia or hypoplastic type); ALX3 (Aristaless-like homeobox. . .

related to AF4); LYLI (Lymphoblastic, leukemia derived sequence 1); MAFG (V-maf musculoaponeurotic fibrosarcoma (avian) oncogene family, protein G); MAX (MAX protein); MDM2 (Mouse double minute 2, human homolog of; p53-binding protein); MHC2TA (MHC class II transactivator); MKI67 (Antigen identified by monoclonal antibody Ki-67); MNDA (Myeloid cell nuclear differentiation antigen); MSXI (Msh (Drosophila) homeo box homolog I (formerly homeo box 7));. . .

integrin-associated signal transducer)); CD5 (CD5 antigen (p56-62)); CD53 (CD53 antigen); CD58 (CD58 antigen, (lymphocyte function-associated antigen 3)); CD59 (CD59 antigen p18-20 (antigen identified by monoclonal antibodies 16.3A5, EJ16, EJ30, EL32 and G344)); CD5L (CD5 antigen-like (scavenger receptor cysteine rich family)); CD6 (CD6 antigen);. . .

LYN (V-src-1 Yamaguchi sarcoma viral related oncogene homolog); LYZ (Lysozyme (renal amyloidosis))-; M1SI (Membrane component, chromosome 1, surface marker I (400 glycoprotein, identified by monoclonal antibody GA733)); MAB21L1 (Mab-21 (C. elegans)-like 1); MACAM1 (Mucosal addressin cell adhesion molecule-1); MADHI (MAD (mothers against decapentaplegic, Drosophila). . . MCC (Mutated in colorectal cancers); MCF2 (MCF.2 cell line derived transforming sequence); MCP (Membrane cofactor protein (CD46, trophoblast-lymphocyte cross-reactive antigen)); MDF1 (Antigen identified by monoclonal antibody A-3A4); MDH2

(Malate dehydrogenase 2, NAD (mitochondrial)); MDUI (Antigen identified by monoclonal antibodies 4172, TRALIO, TROP4, and T43); MEI (Malic enzyme 1, soluble); ME2 (Malic enzyme 2, mitochondrial); MEKKI (MAP/ERK kinase kinase. . . MEMOI (Methylation modifier for class I HLA); MENI (Multiple endocrine neoplasia 1); MEPIA (Meprin A, alpha (PABA peptide hydrolase)); MER2 (Antigen identified by monoclonal antibodies 1D12, 2177); MFAP2 (Microfibrillar-associated protein 2); MFAP4 (Microfibrillar-associated protein 4); MFTS (Migraine, familial typical, susceptibility to); MGCT (MGI); MGP (Matrix Gla protein); MHC2TA (MHC class 11 transactivator); MIC2 (Antigen identified by monoclonal antibodies 12E7, F21 and 013); MIC5 (Antigen identified by monoclonal antibody RI); MIC7 (Antigen identified by monoclonal antibody 28 7); MICA (MHC class I polypeptide-related sequence A); MIF (Macrophage migration inhibitory factor (glycosylation-inhibiting factor)); MIG (Monokine induced. . .  
 .  
 (Uridine phosphorylase); UPK1B (Uroplakin 113); UROD (Uroporphyrinogen decarboxylase); UROS (Uroporphyrinogen III synthase (congenital erythropoietic porphyria)); USH2A (Usher syndrome 2A (autosomal recessive, mild)); USP7 (Ubiquitin specific protease 7 (herpes virus-associated)); VASP (Vasodilator-stimulated phosphoprotein); VCAM 1 (Vascular cell adhesion molecule 1); VDAC 1 (Voltage-dependent anion. . .  
 .  
 CD48 (CD48 antigen (B-cell membrane protein)); CD53 (CD53 antigen); CD58 (CD58 antigen, (lymphocyte function-associated antigen 3)); CD59 (CD59 antigen p18-20 (antigen identified by monoclonal antibodies 16.3A5, EJ16, EJ30, EL32 and G344)); CD63 (CD63 antigen (melanoma 1 antigen)); CD68 (CD68 antigen); CD7 (CD7 antigen. . . gene 3); LY64 (Lymphocyte antigen 64 (mouse) homolog, radioprotective, 105kD); LYZ (Lysozyme (renal amyloidosis)); MAPIB (Microtubule-associated protein 113); MDUI (Antigen identified by monoclonal antibodies 4172, TRALIO, TROP4, and T43); MIC2 (Antigen identified by monoclonal antibodies 12E7, F21 and 013); MICA (MHC class I polypeptide-related sequence A); MME (Membrane metallo-endopeptidase (neutral endopeptidase, enkephalinase, CALLA, . . .  
 melanogaster muscleblind B protein); MDM2 (Mouse double minute 2, human homolog of, p53-binding protein); MHC2TA (MHC class 11 transactivator); MKI67 (Antigen identified by monoclonal antibody Ki-67); MNDA (Myeloid cell nuclear differentiation antigen); MSX1 (Msh (Drosophila) homeo box homolog 1 (formerly homeo box 7)); MTHFD. . .  
 .  
 member 3)); LYN (V-src-1 Yamaguchi sarcoma viral related oncogene homolog); MIS 1 (Membrane component, chromosome 1, surface marker I

(40kD glycoprotein, identified by monoclonal antibody GA733)); M4SI (Membrane component, chromosomal 4, surface marker (35kD glycoprotein)); MADH4 (MAD (mothers against decapentaplegic, Drosophila) homolog. . . oncogene: family, protein K); MASI (MASI oncogene); MAX (MAX protein); MCC (Mutated in colorectal cancers); MCF2 (MCF.2 cell line derived transforming sequence); MDM2 (Mouse double minute 2, human homolog of-, p53-binding protein); MEL (Mel transforming oncogene (derived from cell line NK14)- RAB8 homolog); MELLI (Mel. . . member 1)); LTB (Lymphotoxin beta (TNF superfamily, member 3)); MIS I (Membrane component, chromosome 1, surface marker I (40kD glycoprotein, identified by monoclonal antibody GA733)); M4SI (Membrane component, chromosomal 4, surface marker (35kD glycoprotein)); MADH4. (MAD (mothers against decapentaplegic, Drosophila) homolog 4);. . .

CLMEN. . . according to any of claims I to 4, characterised in that the biological samples A and B are obtained from the identical individual, tissue, cell or other biological material. . Method according claim 5, characterised in that the biological samples A. and B. . .

28 Method according to any of claims I to 27, characterised in that the identical biological sample, different biological samples or a combination thereof is used in steps a) and/or b).

=> d his

(FILE 'HOME' ENTERED AT 15:06:58 ON 20 SEP 2006)

FILE 'CAPLUS' ENTERED AT 15:07:09 ON 20 SEP 2006

L1 21 S HAUSP AND MDM2  
L2 6 S L1 NOT PY>2004  
L3 40 S USP7  
L4 8 S L3 AND MDM2

FILE 'PCTFULL' ENTERED AT 15:12:56 ON 20 SEP 2006

L5 37 S USP7  
L6 34 S HAUSP  
L7 59 S L6 OR L5  
L8 18 S MDM2 AND L7  
L9 532010 S SCREEN? OR IDENT?  
L10 18 S L9 AND L8  
L11 5 S L10 NOT PY>2002

FILE 'REGISTRY' ENTERED AT 15:17:14 ON 20 SEP 2006

L12 1 S E4  
E "HAUSP"/CN 25  
E "USP7"/CN 25  
E "USP 7"/CN 25  
E "USP-7"/CN 25

FILE 'MEDLINE' ENTERED AT 15:19:32 ON 20 SEP 2006

L13 39 S HAUSP OR (USP ( ) 7)  
L14 55 S HAUSP OR (USP7)

L15 2699 S MDM2  
L16 18 S L15 AND L14  
L17 1 S L16 NOT PY>2002

FILE 'PCTFULL' ENTERED AT 15:24:49 ON 20 SEP 2006

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L2 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:312567 CAPLUS  
DOCUMENT NUMBER: 137:44608  
TITLE: Deubiquitination of p53 by HAUSP is an important pathway for p53 stabilization  
AUTHOR(S): Li, Muyang; Chen, Delin; Shiloh, Ariel; Luo, Jianyuan; Nikolaev, Anatoly Y.; Qin, Jun; Gu, Wei  
CORPORATE SOURCE: Institute for Cancer Genetics, and Department of Pathology, College of Physicians b Surgeons, Columbia University, New York, NY, 10032, USA  
SOURCE: Nature (London, United Kingdom) (2002), 416(6881), 648-652  
CODEN: NATUAS; ISSN: 0028-0836  
PUBLISHER: Nature Publishing Group  
DOCUMENT TYPE: Journal  
LANGUAGE: English

AB The p53 tumor suppressor is a short-lived protein that is maintained at low levels in normal cells by Mdm-mediated ubiquitination and subsequent proteolysis. Stabilization of p53 is crucial for its tumor suppressor function. However, the precise mechanism by which ubiquitinated p53 levels are regulated in vivo is not completely understood. By mass spectrometry of affinity-purified p53-associated factors, the authors have identified herpesvirus-associated ubiquitin-specific protease (HAUSP) as a novel p53-interacting protein. HAUSP strongly stabilizes p53 even in the presence of excess Mdm2, and also induces p53-dependent cell growth repression and apoptosis. Significantly, HAUSP has an intrinsic enzymic activity that specifically



deubiquitinates p53 both in vitro and in vivo. In contrast, expression of a catalytically inactive point mutant of HAUSP in cells increases the levels of p53 ubiquitination and destabilizes p53. These findings reveal an important mechanism by which p53 can be stabilized by direct deubiquitination and also imply that HAUSP might function as a tumor suppressor in vivo through the stabilization of p53.

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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